

10/ 774, 415

FILE COVERS 1907 - 12 Mar 2003 VOL 138 ISS 11  
FILE LAST UPDATED: 11 Mar 2003 (20030311/ED)

This file contains CAS Registry Numbers for easy and accurate  
substance identification.

=> s us 6482848/pn  
L1 3 US 6482848/PN  
(US6482848/PN)

=> s l1  
L2 3 US 6482848/PN  
(US6482848/PN)

=> select l1  
ENTER ANSWER NUMBER OR RANGE (1-):3  
ENTER DISPLAY CODE (TI) OR ?:rn  
E1 THROUGH E37 ASSIGNED

=> file reg	SINCE FILE	TOTAL
COST IN U.S. DOLLARS	ENTRY	SESSION
FULL ESTIMATED COST	5.20	5.41

FILE 'REGISTRY' ENTERED AT 11:13:46 ON 12 MAR 2003  
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.  
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.  
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Property values tagged with IC are from the ZIC/VINITI data file  
provided by InfoChem.

STRUCTURE FILE UPDATES: 11 MAR 2003 HIGHEST RN 497913-82-3  
DICTIONARY FILE UPDATES: 11 MAR 2003 HIGHEST RN 497913-82-3

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when  
conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP  
PROPERTIES for more information. See STNote 27, Searching Properties  
in the CAS Registry File, for complete details:  
<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

=> s e1-e31  
1 106-89-8/BI  
(106-89-8/RN)  
1 108-49-6/BI  
(108-49-6/RN)  
1 109-01-3/BI  
(109-01-3/RN)  
1 111-95-5/BI  
(111-95-5/RN)  
1 123-75-1/BI  
(123-75-1/RN)

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10/774, 465

10243942

1 147-85-3/BI  
(147-85-3/RN)  
1 15285-59-3/BI  
(15285-59-3/RN)  
1 1892-57-5/BI  
(1892-57-5/RN)  
1 2199-51-1/BI  
(2199-51-1/RN)  
1 2199-59-9/BI  
(2199-59-9/RN)  
1 253870-02-9/BI  
(253870-02-9/RN)  
1 2917-91-1/BI  
(2917-91-1/RN)  
1 326914-13-0/BI  
(326914-13-0/RN)  
1 356068-86-5/BI  
(356068-86-5/RN)  
1 356068-89-8/BI  
(356068-89-8/RN)  
1 372092-80-3/BI  
(372092-80-3/RN)  
1 375387-20-5/BI  
(375387-20-5/RN)  
1 375798-45-1/BI  
(375798-45-1/RN)  
1 375798-46-2/BI  
(375798-46-2/RN)  
1 375798-47-3/BI  
(375798-47-3/RN)  
1 375798-48-4/BI  
(375798-48-4/RN)  
1 375798-49-5/BI  
(375798-49-5/RN)  
1 375798-50-8/BI  
(375798-50-8/RN)  
1 375798-51-9/BI  
(375798-51-9/RN)  
1 375798-52-0/BI  
(375798-52-0/RN)  
1 375798-53-1/BI  
(375798-53-1/RN)  
1 375798-54-2/BI  
(375798-54-2/RN)  
1 375798-55-3/BI  
(375798-55-3/RN)  
1 443-69-6/BI  
(443-69-6/RN)  
1 498-63-5/BI  
(498-63-5/RN)  
1 56341-41-4/BI  
(56341-41-4/RN)

L3

31 (106-89-8/BI OR 108-49-6/BI OR 109-01-3/BI OR 111-95-5/BI OR  
123-75-1/BI OR 147-85-3/BI OR 15285-59-3/BI OR 1892-57-5/BI OR  
2199-51-1/BI OR 2199-59-9/BI OR 253870-02-9/BI OR 2917-91-1/BI  
OR 326914-13-0/BI OR 356068-86-5/BI OR 356068-89-8/BI OR 372092-  
80-3/BI OR 375387-20-5/BI OR 375798-45-1/BI OR 375798-46-2/BI  
OR 375798-47-3/BI OR 375798-48-4/BI OR 375798-49-5/BI OR 375798-  
50-8/BI OR 375798-51-9/BI OR 375798-52-0/BI OR 375798-53-1/BI  
OR 375798-54-2/BI OR 375798-55-3/BI OR 443-69-6/BI OR 498-63-5/BI

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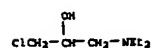
I OR 56341-41-4/BI)

=> d scan

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LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IN 2-Propanol, 1-chloro-1-(diethylamino)- (6CI, 7CI, 8CI, 9CI)  
 MP C7 H16 Cl N O  
 CI COM

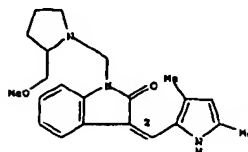


\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

HOW MANY MORE ANSWERS DO YOU WISH TO SCAN? (1):30

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IN 2H-indol-2-one, 3-[(2,5-dimethyl-1H-pyrrol-2-yl)methylene]-1,2-dihydro-1-  
 [(2-(methoxymethyl)-1-pyrrolidinyl)methyl]-, (3Z)- (9CI)  
 MP C12 H17 N3 O2

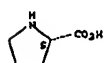
Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

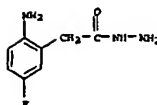
LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IN L-Proline (9CI)  
 MP C5 H9 N O2  
 CI COM

Absolute stereochemistry. Rotation (-).



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IN Benzenecarboxic acid, 2-amino-5-fluoro-, hydrazide (9CI)  
 MP C8 H10 F N3 O

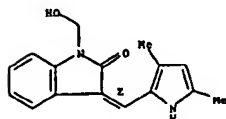


\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

10243942

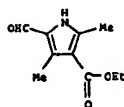
LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN 2H-Indol-2-one, 3-((3,5-dimethyl-1H-pyrrol-2-yl)methylene)-1,3-dihydro-1-  
(hydroxymethyl)-, (3Z)- (9CI)  
MF C16 H16 N2 O3  
CI 6

Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

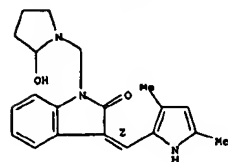
LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN 1H-Pyrrole-3-carboxylic acid, 5-formyl-2,4-dimethyl-, ethyl ester (9CI)  
MF C10 H13 N O3  
CI 6



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN 2H-Indol-2-one, 3-((1,5-dimethyl-1H-pyrrol-2-yl)methylene)-1,3-dihydro-1-  
((2-hydroxy-1-pyrrolidinyl)methyl)-, (3Z)- (9CI)  
MF C20 H23 N3 O2  
CI 6

Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

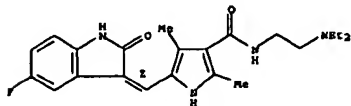
LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN Ethanamine, 2-methoxy-N-(2-methoxyethyl)- (9CI)  
MF C6 H15 N O2  
CI 6

MeO-CH2-CH2-NH-CH2-CH2-OMe

\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IN 1H-Pyrrole-3-carboxamide, N-(2-(diethylamino)ethyl)-5-[(E)-(5-fluoro-1,3-dihydro-2-oxo-1H-indol-1-ylidene)methyl]-2,4-dimethyl- (9CI)  
 MF C22 H27 F N4 O2  
 CI COM

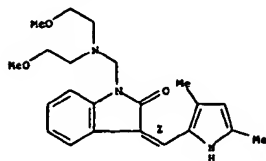
Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IN 2H-Indol-3-one, 1-[(bis(2-methoxyethyl)amino)methyl]-3-[(1,5-dimethyl-1H-pyrrol-2-yl)methylene]-1,3-dihydro-, (3Z)- (9CI)  
 MF C22 H29 N3 O3  
 CI COM

Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

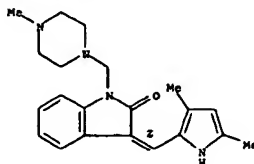
LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IN 1,3-Propanediamine, N'-(ethylcarbonimidoyl)-N,N-dimethyl- (9CI)  
 MF C8 H17 N3  
 CI COM

$\text{Et}-\text{N}=\text{C}=\text{N}-(\text{CH}_2)_3-\text{NMe}_2$

\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IN 2H-Indol-2-one, 3-[(1,5-dimethyl-1H-pyrrol-2-yl)methylene]-1,3-dihydro-1-[(4-methyl-1-piperazinyl)methyl]-, (3Z)- (9CI)  
 MF C21 H26 N4 O  
 CI COM

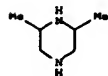
Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

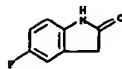
10243942

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN Piperazine, 2,6-dimethyl- (7CI, 8CI, 9CI)  
MF C6 H14 N2  
CI COM



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

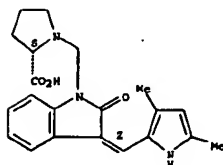
LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN 2H-Indol-2-one, 5-fluoro-1,3-dihydro- (9CI)  
MF C8 H6 F N O



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

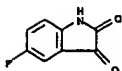
LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN L-Proline, 1-[[[(3E)-3-[(2,6-dimethyl-1H-pyrrol-2-yl)methylene]-2,3-dihydro-2-oxo-1H-indol-1-yl)methyl]- (9CI)  
MF C21 H23 N3 O3

Absolute stereochemistry.  
Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN 1H-Indole-2,3-dione, 5-fluoro- (9CI)  
MF C8 H4 F N O2



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

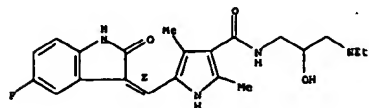
10243942

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IN Kinase (phosphorylating), protein (9CI)  
 MP Unspecified  
 CI MAH

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

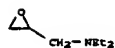
LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IN 1H-Pyrrole-3-carboxamide, N-[3-(diethylamino)-2-hydroxypropyl]-5-[[2]-[5-fluoro-1,2-dihydro-3-oxo-1H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI)  
 MP C3 H29 F N4 O3

Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

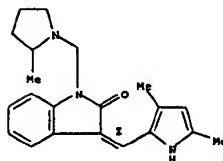
LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IN Oxiranemethanamine, N,N-diethyl- (9CI)  
 MP C7 H15 N O  
 CI COM



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IN 2H-Indol-2-one, 3-[[[1,5-dimethyl-1H-pyrrol-2-yl)methylene]-1,3-dihydro-1-[[2-methyl-1-pyrrolidinyl)methyl]-, (3Z)- (9CI)  
 MP C21 H25 N3 O

Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*



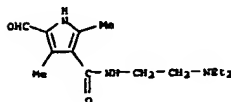
10243942

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IM Pyrrolidine (8CI, 9CI)  
 MP C4 H9 N  
 CI COM, RPS



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

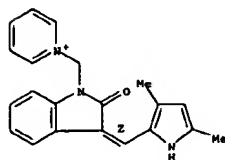
LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IM 1H-Pyrrole-3-carboxamide,  
 N-[2-(diethylamino)ethyl]-5-formyl-2,4-dimethyl-  
 (9CI)  
 MP C14 H23 N3 O2  
 CI



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

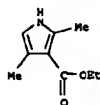
LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IM Pyridinium, 1-[[[1,3,5-trimethyl-1H-pyrrol-2-yl)methylene]-2,3-  
 dihydro-2-oxo-1H-indol-1-yl)methyl]-, chloride (9CI)  
 MP C21 H20 N3 O . Cl  
 CI

Double bond geometry as shown.



• Cl<sup>-</sup>

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IM 1H-Pyrrole-3-carboxylic acid, 2,4-dimethyl-, ethyl ester (9CI)  
 MP C9 H13 N O2  
 CI COM

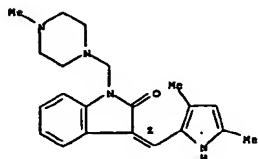


\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

10243942

L3 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN 2H-Indol-2-one, 1-[(2,5-dimethyl-1H-pyrrol-2-yl)methylene]-1,3-dihydro-1-  
[(4-methyl-1-piperazinyl)methyl]-, dihydrochloride, (2Z)- (9CI)  
MF C21 H26 N4 O . 3 Cl H

Double bond geometry as shown.



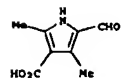
• 2 HCl

L3 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN Piperazine, 1-methyl- (8CI, 9CI)  
MF C5 H12 N2  
CI COM



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

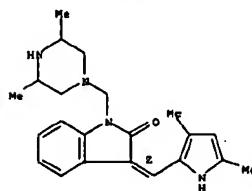
L3 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN 1H-Pyrrole-2-carboxylic acid, 5-formyl-2,4-dimethyl- (9CI)  
MF C6 H8 N O3



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

L3 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN 2H-Indol-2-one,  
1-[(3,5-dimethyl-1-piperazinyl)methyl]-1-[(2,5-dimethyl-1H-  
pyrrol-2-yl)methylene]-1,3-dihydro-, (2Z)- (9CI)  
MF C22 H28 N4 O

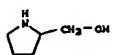
Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

10243942

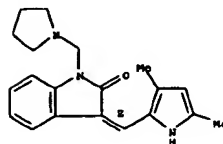
LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IM 2-Pyrrolidinomethanol (6CI, 7CI, 8CI, 9CI)  
 MF C5 H11 N O  
 CI COM



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IM 28-Indol-3-one, 3-[[[1,3-dimethyl-4H-pyrrol-2-yl)methylene]-1,1-dihydro-1H-pyrrolidin-2-ylidene]-, (3E)- (9CI)  
 MF C10 H13 N3 O  
 CI COM

Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

LJ 31 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
 IM Oxirane, (chloromethyl)- (9CI)  
 MF C2 H5 Cl O  
 CI COM



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

ALL ANSWERS HAVE BEEN SCANNED

=> logoff

ALL L# QUERIES AND ANSWER SETS ARE DELETED AT LOGOFF

LOGOFF? (Y)/N/HOLD:y

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

0.80

6.21

STN INTERNATIONAL LOGOFF AT 11:14:55 ON 12 MAR 2003

10/724, 415

~~10243365~~

ENERGY, INSPEC  
NEWS 43 Feb 13 CANCERLIT is no longer being updated  
NEWS 44 Feb 24 METADEX enhancements  
NEWS 45 Feb 24 PCTGEN now available on STN  
NEWS 46 Feb 24 TBMA now available on STN  
NEWS 47 Feb 26 NTIS now allows simultaneous left and right truncation  
NEWS 48 Feb 26 PCTFULL now contains images  
NEWS 49 Mar 04 SDI PACKAGE for monthly delivery of multifile SDI results

NEWS EXPRESS January 6 CURRENT WINDOWS VERSION IS V6.01a,  
CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),  
AND CURRENT DISCOVER FILE IS DATED 01 OCTOBER 2002  
NEWS HOURS STN Operating Hours Plus Help Desk Availability  
NEWS INTER General Internet Information  
NEWS LOGIN Welcome Banner and News Items  
NEWS PHONE Direct Dial and Telecommunication Network Access to STN  
NEWS WWW CAS World Wide Web Site (general information)

Enter NEWS followed by the item number or name to see news on that specific topic.

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\* \* \* \* \* STN Columbus \* \* \* \* \*

FILE 'HOME' ENTERED AT 11:27:04 ON 12 MAR 2003

=> FILE REG  
COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
0.21	0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 11:27:10 ON 12 MAR 2003  
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.  
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.  
COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 11 MAR 2003 HIGHEST RN 497913-82-3  
DICTIONARY FILE UPDATES: 11 MAR 2003 HIGHEST RN 497913-82-3

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details:

Kamal Saeed

10243942

<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

=> ....Testing the current file.... screen

ENTER SCREEN EXPRESSION OR (END):end

=>

Uploading C:\Program Files\Stnexp\Queries\10243942.str

L1       STRUCTURE UPLOADED

=> que L1

L2    QUE L1

=> S L1 FULL

FULL SEARCH INITIATED 11:33:33 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED -   3431 TO ITERATE

100.0% PROCESSED       3431 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

L3               0 SEA SSS FUL L1

=> ....Testing the current file.... screen

ENTER SCREEN EXPRESSION OR (END):end

=>

Uploading C:\Program Files\Stnexp\Queries\10243942.str

L4       STRUCTURE UPLOADED

=> que L4

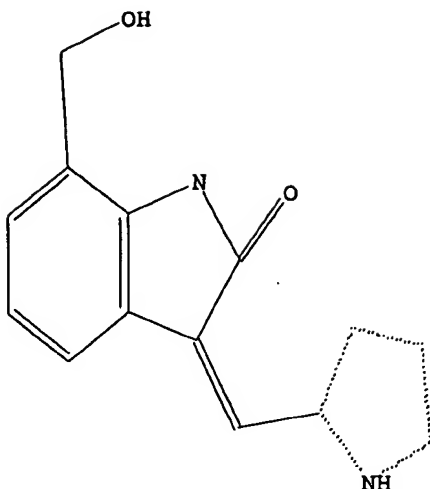
L5    QUE L4

=> D

L5 HAS NO ANSWERS

L4               STR

10243942



Structure attributes must be viewed using STN Express query preparation.  
L5                    QUE ABB=ON PLU=ON L4

=> S L4 FULL  
FULL SEARCH INITIATED 11:34:53 FILE 'REGISTRY'  
FULL SCREEN SEARCH COMPLETED - 3431 TO ITERATE

100.0% PROCESSED      3431 ITERATIONS                    0 ANSWERS  
SEARCH TIME: 00.00.01

L6                    0 SEA SSS FUL L4

=> ....Testing the current file.... screen

ENTER SCREEN EXPRESSION OR (END):end

=>  
Uploading C:\Program Files\Stnexp\Queries\10243942.str

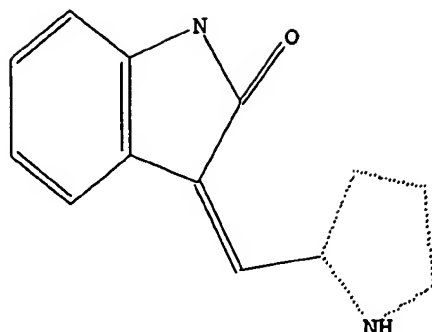
L7                    STRUCTURE UPLOADED

=> que L7

L8                    QUE L7

=> D  
L8 HAS NO ANSWERS  
L7                    STR

10243942



Structure attributes must be viewed using STN Express query preparation.  
L8 QUE ABB=ON PLU=ON L7

=> S L7 FULL  
FULL SEARCH INITIATED 11:36:40 FILE 'REGISTRY'  
FULL SCREEN SEARCH COMPLETED - 5268 TO ITERATE

100.0% PROCESSED 5268 ITERATIONS 1860 ANSWERS  
SEARCH TIME: 00.00.01

L9 1860 SEA SSS FUL L7

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	449.65	449.86

FILE 'CAPLUS' ENTERED AT 11:36:51 ON 12 MAR 2003  
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FILE COVERS 1907 - 12 Mar 2003 VOL 138 ISS 11  
FILE LAST UPDATED: 11 Mar 2003 (20030311/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L9  
L10 128 L9

Kamal Saeed



10243942

=> FILE REG  
COST IN U.S. DOLLARS

SINCE FILE ENTRY	TOTAL SESSION
0.83	450.69

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 11:38:08 ON 12 MAR 2003  
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provided by InfoChem.

STRUCTURE FILE UPDATES: 11 MAR 2003 HIGHEST RN 497913-82-3  
DICTIONARY FILE UPDATES: 11 MAR 2003 HIGHEST RN 497913-82-3

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when  
conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP  
PROPERTIES for more information. See STNote 27, Searching Properties  
in the CAS Registry File, for complete details:  
<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

=> ....Testing the current file.... screen

ENTER SCREEN EXPRESSION OR (END):end

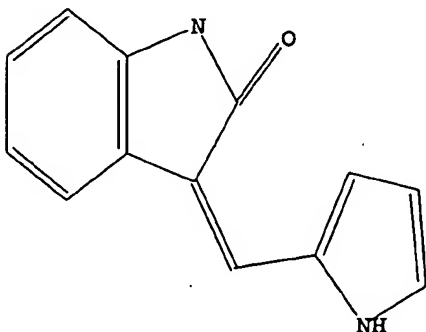
=>  
Uploading C:\Program Files\Stnexp\Queries\10243942.str

L11 STRUCTURE UPLOADED

=> que L11

L12 QUE L11

=> D  
L12 HAS NO ANSWERS  
L11 STR



Kamal Saeed

10243942

Structure attributes must be viewed using STN Express query preparation.  
L12                    QUE ABB=ON PLU=ON L11

=> S L11 FULL  
FULL SEARCH INITIATED 11:38:31 FILE 'REGISTRY'  
FULL SCREEN SEARCH COMPLETED - 3074 TO ITERATE

100.0% PROCESSED      3074 ITERATIONS                    1860 ANSWERS  
SEARCH TIME: 00.00.01

L13                    1860 SEA SSS FUL L11

=> FILE REG		
COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	148.55	599.24

FILE 'REGISTRY' ENTERED AT 11:39:18 ON 12 MAR 2003  
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.  
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STRUCTURE FILE UPDATES: 11 MAR 2003 HIGHEST RN 497913-82-3  
DICTIONARY FILE UPDATES: 11 MAR 2003 HIGHEST RN 497913-82-3

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when  
conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP  
PROPERTIES for more information. See STNote 27, Searching Properties  
in the CAS Registry File, for complete details:  
<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

=> ....Testing the current file.... screen

ENTER SCREEN EXPRESSION OR (END):end

=>  
Uploading C:\Program Files\Stnexp\Queries\10243942.str

L14                    STRUCTURE UPLOADED

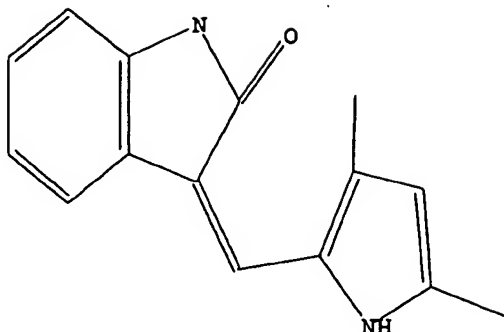
=> que L14

L15                    QUE L14

=> D  
L15 HAS NO ANSWERS  
L14                    STR

Kamal Saeed

10243942



Structure attributes must be viewed using STN Express query preparation.  
L15 QUE ABB=ON PLU=ON L14

=> S L14 FULL  
FULL SEARCH INITIATED 11:39:40 FILE 'REGISTRY'  
FULL SCREEN SEARCH COMPLETED - 2980 TO ITERATE

100.0% PROCESSED 2980 ITERATIONS 1282 ANSWERS  
SEARCH TIME: 00.00.01

L16 1282 SEA SSS FUL L14

=> FILE CAPLUS		
COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	148.15	747.39

FILE 'CAPLUS' ENTERED AT 11:39:51 ON 12 MAR 2003  
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FILE COVERS 1907 - 12 Mar 2003 VOL 138 ISS 11  
FILE LAST UPDATED: 11 Mar 2003 (20030311/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L14  
REGISTRY INITIATED  
Substance data SEARCH and crossover from CAS REGISTRY in progress...

Kamal Saeed

10243942

Use DISPLAY HITSTR (or FHITSTR) to directly view retrieved structures.

SAMPLE SEARCH INITIATED 11:40:00 FILE 'REGISTRY'  
SAMPLE SCREEN SEARCH COMPLETED - 134 TO ITERATE

100.0% PROCESSED 134 ITERATIONS 50 ANSWERS  
INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)  
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*  
BATCH \*\*COMPLETE\*\*  
PROJECTED ITERATIONS: 1986 TO 3374  
PROJECTED ANSWERS: 720 TO 1640

L17 50 SEA SSS SAM L14

L18 8 L17

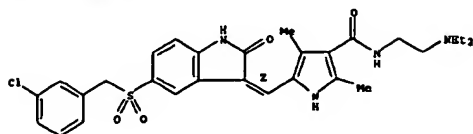
=> D IBIB ABS HITSTR TOT



10243942

L16 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)  
 dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(diethylamino)ethyl]-3,4-  
 dimethyl- (9CI) (CA INDEX NAME)

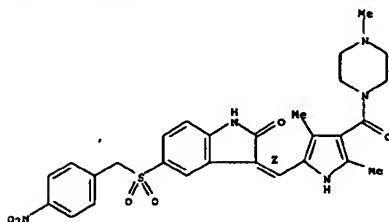
Double bond geometry as shown.



RN 477574-10-0 CAPLUS

CN Piperazine,  
 1-[[5-[(2)-[1,2-dihydro-5-[[[4-nitrophenyl)methyl]sulfonyl]-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]-4-methyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

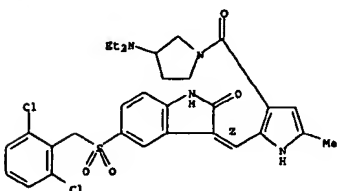


RN 477574-45-1 CAPLUS

CN Piperazine,  
 1-[[5-[(2)-[1,2-dihydro-5-[[[2,6-difluorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]-4-methyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

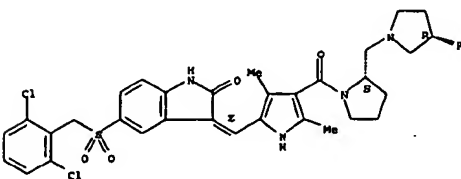
L16 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



RN 477575-04-5 CAPLUS

CN Pyrrolidine, 1-[[5-[(2)-[1,2-dihydro-5-[[[2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]-2-[[[3R]-3-fluoro-1-pyrrolidinyl)methyl]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.  
 Double bond geometry as shown.

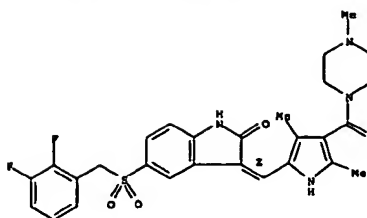


RN 477575-29-4 CAPLUS

CN Piperazine, 1-[[5-[(2)-[1,2-dihydro-5-[[[2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]-4-methyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

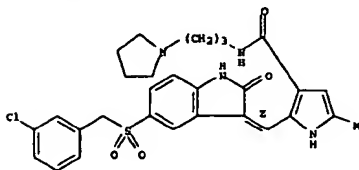
L16 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



RN 477574-75-7 CAPLUS

CN 1H-Pyrrole-3-carboxamide,  
 2-[[5-[(2)-[1,2-dihydro-5-[[[2-chlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-5-methyl-N-[3-[[1-pyrrolidinyl]propyl]- (9CI) (CA INDEX NAME)

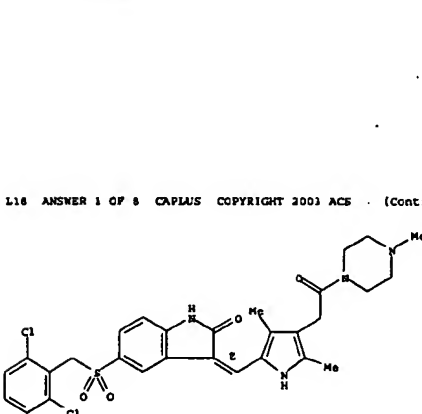
Double bond geometry as shown.



RN 477574-96-2 CAPLUS

CN 3-Pyrrolidinamine, 1-[[5-[(2)-[1,2-dihydro-5-[[[2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-5-methyl-N-[3-[[1-pyrrolidinyl]propyl]- (9CI) (CA INDEX NAME)

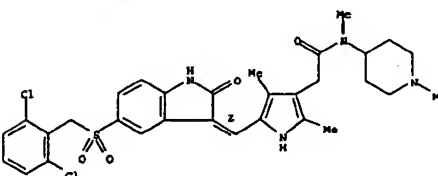
Double bond geometry as shown.



RN 477575-32-9 CAPLUS

CN 1H-Pyrrole-3-acetamide, 5-[[5-[(2)-[1,2-dihydro-5-[[[2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N,2,4-trimethyl-N-(1-methyl-4-piperidinyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.



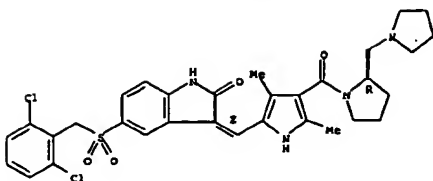
RN 477575-56-7 CAPLUS

CN Pyrrolidine, 1-[[5-[(2)-[1,2-dihydro-5-[[[2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]-2-[[1-pyrrolidinyl)methyl]-, (2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.  
 Double bond geometry as shown.

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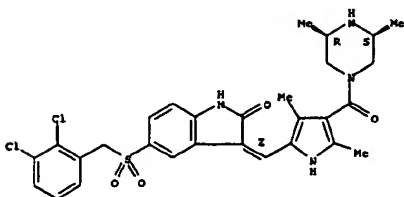
L18 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



RN 477575-69-2 CAPLUS

CN Piperazine, 1-([5-[(2)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl)-2,5-dimethyl-, (3R,5S)-(9CI) (CA INDEX NAME)

Absolute stereochemistry.  
Double bond geometry as shown.

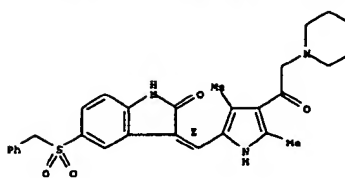


RN 477575-88-5 CAPLUS

CN 3H-Indol-2-one, 3-([3,5-dimethyl-4-[(1-piperidinyl)acetyl]-1H-pyrrol-2-yl]methylene)-1,3-dihydro-5-[(phenylmethyl)sulfonyl]-, (3Z)-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

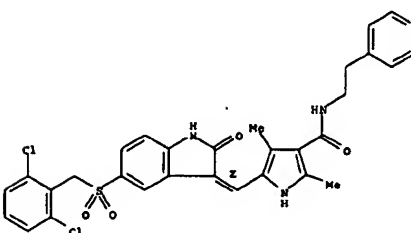
L18 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



RN 477575-90-3 CAPLUS

CN 1H-Pyrrole-3-carboxamide, 5-([(2)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[2-(6-pyridinyl)ethyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

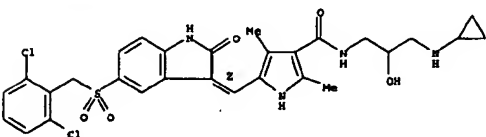


RN 477576-52-6 CAPLUS

CN 1H-Pyrrole-3-carboxamide, N-[3-(cyclopropylamino)-2-hydroxypropyl]-5-([(2)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

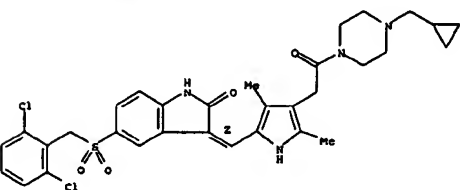
L18 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



RN 477576-65-1 CAPLUS

CN Piperazine, 1-(cyclopropylmethyl)-4-([(5-[(2)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]acetyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

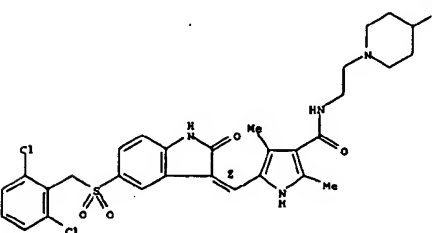


RN 477576-95-7 CAPLUS

CN 1H-Pyrrole-3-carboxamide, 5-([(2)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-[2-(4-fluoro-1-piperidinyl)ethyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

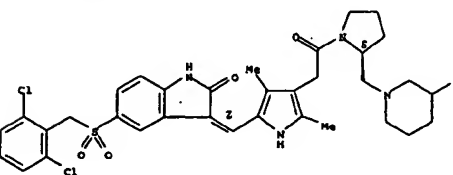
L18 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



RN 477577-09-6 CAPLUS

CN Pyrrolidine, 1-([(5-[(2)-[5-[(2,6-dichlorophenyl)methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]acetyl)-2-[(3-fluoro-1-piperidinyl)methyl]-, (2S)-(9CI) (CA INDEX NAME)

Absolute stereochemistry.  
Double bond geometry as shown.



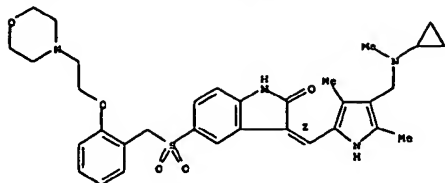
RN 477577-54-1 CAPLUS

CN 3H-Indol-2-one, 3-([4-[(cyclopropylmethylanino)methyl]-3,5-dimethyl-1H-pyrrol-2-yl]methylene)-1,3-dihydro-5-([(2)-[2-(4-morpholinyl)ethoxy]phenyl)methyl]sulfonyl)-, (3Z)-(9CI) (CA INDEX NAME)

Double bond geometry as shown.

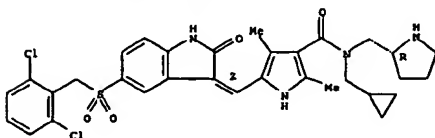
10243942

L18 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



RN 477577-66-5 CAPLUS  
 CN 1H-Pyrrole-3-carboxamide, N-(cyclopropylmethyl)-5-((Z)-[5-[[[2,6-dichlorophenyl]methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[(2R)-2-pyrrolidinylmethyl]- (9CI) (CA INDEX NAME)

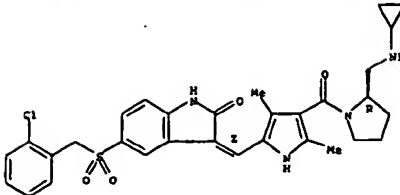
Absolute stereochemistry.  
 Double bond geometry as shown.



RN 477577-74-5 CAPLUS  
 CN 2-Pyrrolidinemethanamine, 1-[[[5-((Z)-[5-[[[2-chlorophenyl]methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]-N-cyclopropyl-, (2R)- (9CI) (CA INDEX NAME)

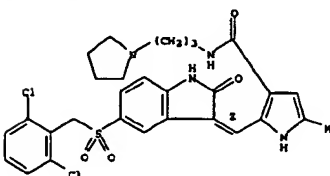
Absolute stereochemistry.  
 Double bond geometry as shown.

L18 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



RN 477578-02-3 CAPLUS  
 CN 1H-Pyrrole-3-carboxamide, 2-((Z)-[5-[[[2,6-dichlorophenyl]methyl]sulfonyl]-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-5-methyl-N-[(3-[(1-pyrrolidinyl)propyl]- (9CI) (CA INDEX NAME)

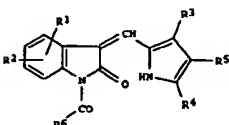
Double bond geometry as shown.



L18 ANSWER 2 OF 8 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002792619 CAPLUS  
 DOCUMENT NUMBER: 137.294870  
 TITLE: Preparation of prodrugs of 3-(pyrrol-2-ylmethylidene)-2-indolinones and activity as modulators of protein kinases  
 INVENTOR(S): Sun, Connie Li; Wei, Chung Chen; Tang, Peng Cho; Koenig, Marcel; Zhou, Yong; Vojtkovsky, Tomas; Nemethy, Adam S.  
 PATENT ASSIGNER(S): Sugen, Inc., USA  
 SOURCE: PCT Int. Appl., 194 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002081466	A1	20021017	MO 2002-US11001	20020409
<p>AB, AD, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BE, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DO, EC, EE, ES, FI, GB, GD, GE, GR, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MY, NZ, OM, PA, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AE, BT, KG, KZ, MD, RU, TJ, TN</p> <p>RN: CH, CN, DE, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TO</p> <p>PRIORITY APPL. INFO.: US 2001-282630P P 20010409</p> <p>OTHER SOURCE(S): MARPAT 137.294870</p> <p>GI</p>				



AB The present invention relates to pyrrole substituted 2-indolinone compds. (shown as 1; e.g. 3-[(1-(3,5-dimethyl-1H-pyrrol-2-yl)methyl-(Z)-ylidene]-2-oxo-2,3-dihydroindole-1-carboxyl chloride) and their pharmaceutically acceptable salts which modulate the activity of protein kinases and therefore are expected to be useful in the prevention and treatment of protein kinase related cellular disorders such as cancer (no data). In

1, R1 and R2 are independently H, halo, alkyl, alkylthio, nitro, trihalomethyl, hydroxy, hydroxyalkyl, alkoxy, cyano, aryl, heteroaryl,

L18 ANSWER 2 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)

C(O)R7 (R7 is alkyl, amino, hydroxy, alkoxy, aryl, heteroaryl, aryloxy, heteroaryloxy, heterocycle, and aminoalkylamino), -NR8R9, -NR8C(O)R9, -SO2R8, and -S(O)2NR8R9 (R8 and R9 are independently H, alkyl, aryl and heteroaryl, or R8 and R9 together with the N to which they are attached form a satd. heterocycloamino). R3 is H, alkyl, hydroxyalkyl, aminoalkyl, -C(O)R7, aryl, and heteroaryl; R4 is H, alkyl, -C(O)R7 aryl, and heteroaryl. R5 is H and -COR10 where R10 is alkyl, alkoxy, hydroxy, aryl, aryloxy, heteroaryl, heterocycle, alkylamino, dialkylamino, or -NR11R12 where R11 is H or alkyl, and R12 is aminoalkyl, hydroxyalkyl, acetylalkyl, cyanoalkyl, carboxyalkyl, alkoxyalkyl, heteroalkyl, or heterocyclylalkyl wherein the alkyl chain in aminoalkyl, heteroalkyl, heteroaryloxy, or heterocyclylalkyl is optionally substituted with one or two hydroxy group(s); or R4 and R5 together form - (CH2)4- or -(CH2)mC(O)CH2- wherein n is 0 to 3, provided that n is 3. R5 is: (c) -OR13 wherein R13 is alkyl, trifluoromethyl, carboxyalkyl, aminoalkyl, phosphonoalkyl, sulfoalkyl, hydroxyalkyl, alkoxyalkyl, aryl, heteroaryl, heteroalkyl, heterocyclyl, monosaccharides and heterocyclylalkyl wherein the alkyl chain in carboxyalkyl, aminoalkyl, phosphonoalkyl, sulfoalkyl, heteroalkyl, heterocyclylalkyl, hydroxyalkyl, or alkoxyalkyl is optionally substituted with one or two hydroxy group(s) and further wherein one or two C atoms in said alkyl chain are optionally replaced by O, -NR14- (R14 is H or alkyl), -S-, or -SO2-; or (d) -NR15R16 where R15 and R16 are independently H, alkyl, carboxyalkyl, alkoxyalkyl, aminoalkyl, phosphonoalkyl, sulfoalkyl, hydroxyalkyl, aryl, heteroaryl, heteroalkyl, and heterocyclylalkyl, wherein the alkyl chain in carboxyalkyl, aminoalkyl, phosphonoalkyl, heteroalkyl, heterocyclylalkyl, hydroxyalkyl, or alkoxyalkyl is optionally substituted with one or two hydroxy group(s) and further wherein one or two C atoms in the alkyl chain are optionally replaced by O, -NR17- (R17 is H or alkyl), -S-, or -SO2-; or R15 and R16 together with

the N atom to which they are attached form satd. or unsatd. heterocycloamino. Although the methods of prep. are not claimed, >80 example preps. are included, both of 1 and the unprotected version of 1 in which the C(O)R6 group has been replaced by H.

17 468744-92-59, 3-[(1-(3,5-dimethyl-1H-pyrrol-2-yl)methyl-(Z)-ylidene)-2-oxo-2,3-dihydroindole-1-carboxylic acid methyl ester  
 RL: PAC (Pharmacological activity); SPW (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

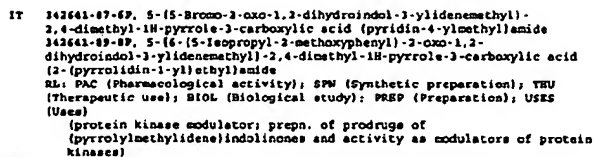
(protein kinase modulator prodrug; prep. of prodrugs of [pyrrol-2-ylmethylidene]indolinones and activity as modulators of protein kinases)

RN 468744-92-5 CAPLUS  
 CN 1H-Indole-1-carboxylic acid, 3-[(1-(3,5-dimethyl-1H-pyrrol-2-yl)methylidene)-2,3-dihydro-2-oxo-, methyl ester, (Z)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.



110 ANSWER 2 OF 8 CAPLUM COPYRIGHT 2003 ACS (Continued)

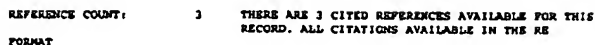


RN 142641-89-8 CAPLUS  
CN 1H-Pyrrole-3-carboxamide, 5-({[1,2-dihydro-6-[2-methoxy-5-{1-methylethyl}phenyl]-2-oxo-3H-indol-3-ylidene]methyl}-2,4-dimethyl-N-[2-(1-pyrrolyldinyl)ethyl]-9CI) (CA INDEX NAME)

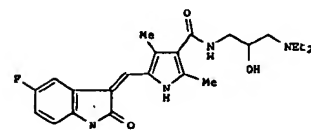
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L18 ANSWER 3 OF 8 CAPLUS COPYRIGHT 2003 ACS
ACCESSION NUMBER: 2002:658111 CAPLUS
DOCUMENT NUMBER: 137:185408
TITLE: 3-(4-Amidopyrrol-2-ylmethylidene)-2-indolinone
        derivatives as protein kinase inhibitors
INVENTOR(S): Guan, Huiping; Liang, Congxin; Sun, Li; Tang, Peng;
              Chao, Wei, Chung Chen; Nauragis, Michael A.;
              Vojtkovsky,
              Tomasz; Jin, Qingwu; Harrington, Paul Matthew
PATENT ASSIGNEE(S): USA
SOURCE: PCT Int. Appl., 167 pp.
        CODEN: PIXX03
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

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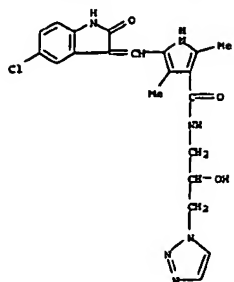
[illegible]

L18 ANSWER 3 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



AB Title compds. [R1 = H, halo, alkyl, haloalkoxy, cycloalkyl, heterocyclic, OH, alkoxy, (un)esterified CO2H, (un)substituted NH2, CONH2];  
R2 = H, halo, alkyl, trihalomethyl, OH, alkoxy, CN, (un)substituted NH2, SO2NH2, (un)esterified CO2H, SO2R8, R8 = alkyl, aryl, aralkyl, heteroaryl, heteroalkyl; R3-R6 = H, alkyl; R7 = H, alkyl, aryl, heteroaryl, acyl, S = aryl, heteroaryl, heterocyclic, (un)substituted NH2] were pred. for use as protein kinase inhibitors in treatment of diseases, such as cancer (no data). Thus, R<sup>1</sup>, 3,5-dimethyl-4-pyridylcarboxylate was oxidized to the 5-carboxaldehyde, followed by ester hydrolysis, reaction with 5-fluoro-2-oxindole and amidation to give the acid 11.  
IT 452104-49-49 452104-99-49  
RL SPW (Synthetic preparation); THU (Therapeutic use); BLOL (Biological study); PREP (Preparation); USES (Uses)  
(prepn. of 3-(4-(amidopyrrol-2-yl)methylidene)-2-indolinone derive. as protein kinase inhibitors)  
RN 452104-49-3 CAPLUS  
CN 1H-Pyrrole-3-carboxamide, 5-[(5-chloro-1,2-dihydro-3-oxo-1H-indol-3-ylidene)methyl]-N-[2-hydroxy-3-[(1H-3,2,3-triazol-1-yl)propyl]-2,4-dimethyl-1H-1,2,4-triazol-5-yl] (CA, IUPAC, NAME)

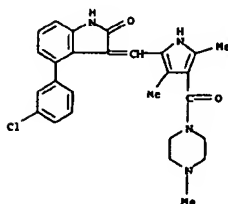
L10 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2003 ACS (Continued)

Cc1c(C(=O)NCC(O)CN2C=NC=C2)c3c(c1)nc4c3c[nH]4C(=O)c5ccc(Br)cc5

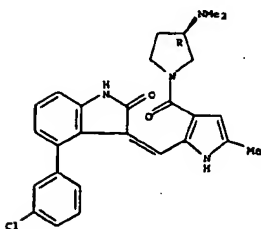
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L10 ANSWER 4 OF 20 CAPLUS COPYRIGHT 2003 ACS (Continued)
442561-58-2P 442561-87-7P 442561-89-9P
442562-68-3P
RW R01 1.00 (Pharmacological activity); SPW (Synthetic preparation); THU
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
(Uses)
(target compd.; prepn. of (eryl)(pyrrolylmethylene)indolinones as
protein kinase signal transduction modulators)
RW 442558-30-7 CAPLUS
CA Piperazine, 1-[5-[(4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-
ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]-4-methyl- (SCI)
INDEX NAME)

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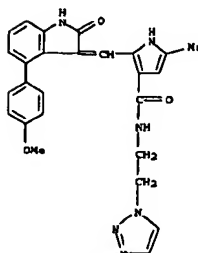
Absolute stereochemistry.  
Double bond geometry unknown.



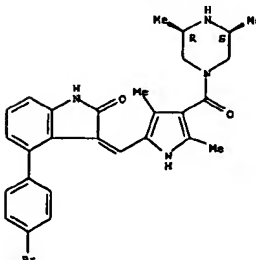
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002055517	A2	20020718	WO 2001-US48564	20011220
WO 2002055517	A3	20020926		

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

L18 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)  
 RN 442568-44-3 CAPLUS  
 CN 1H-Pyrrolo[3-carboxamide, 2-[[[1,2-dihydro-4-(4-methoxyphenyl)-2-oxo-3H-indol-3-ylidene]methyl]-5-methyl-N-[2-(1H-1,2,3-triazol-1-yl)ethyl]-  
 (9CI)  
 (CA INDEX NAME)



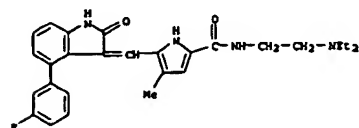
Relative stereochemistry.  
Double bond geometry unknown.



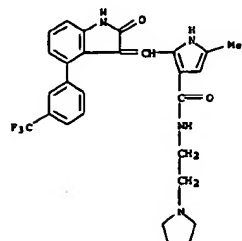
RM 442554-73-0 CAPLUS

10243942

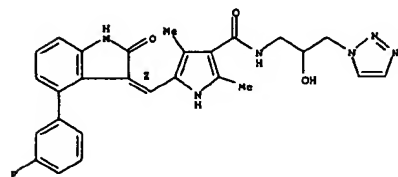
L18 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)  
 RM 1H-Pyrrole-2-carboxamide.  
 CW N-[(2-diethylamino)ethyl]-5-[[4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-4-methyl- (9CI) (CA INDEX NAME)



RM 442559-11-7 CAPLUS  
 CW 1H-Pyrrole-3-carboxamide, 2-[[1,2-dihydro-2-oxo-4-(3-(trifluoromethyl)phenyl)-3H-indol-3-ylidene)methyl]-5-methyl-N-(2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX NAME)

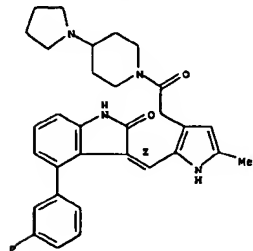


RM 442559-21-9 CAPLUS  
 CW 1H-Pyrrole-3-carboxamide, 2-[[4-(2-chloro-4-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-5-methyl-N-(2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX NAME)



RM 442559-65-1 CAPLUS  
 CW Piperidine, 1-[[2-[[2-[[4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-5-methyl-1H-pyrrol-3-yl]acetyl]-4-(1-pyrrolidinyl)- (9CI) (CA INDEX NAME)

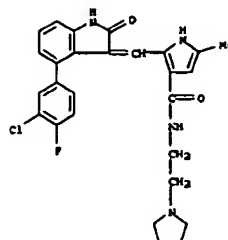
Double bond geometry as shown.



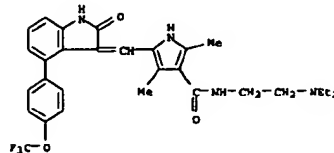
RM 442561-26-4 CAPLUS  
 CW Pyrrolidine, 1-[[2-[[2-[[4-(2-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-6-methyl-1H-pyrrol-3-yl]carbonyl]-2-[[3R]-3-hydroxy-1-pyrrolidinyl)methyl]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.  
 Double bond geometry as shown.

L18 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



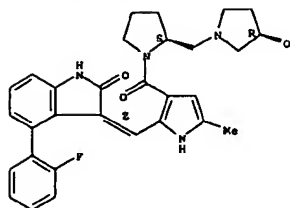
RM 442559-42-4 CAPLUS  
 CW 1H-Pyrrole-3-carboxamide, N-[(2-diethylamino)ethyl]-5-[[1,2-dihydro-2-oxo-4-(4-(trifluoromethoxy)phenyl)-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)



RM 442559-57-1 CAPLUS  
 CW 1H-Pyrrole-3-carboxamide, 5-[(E)-[4-(3-fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-(2-hydroxy-2-(1H-1,2,3-triazol-1-yl)propyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

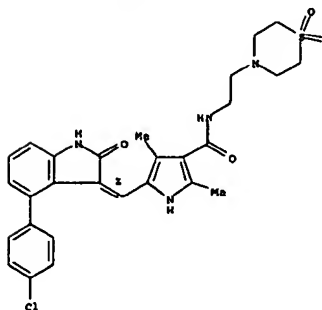
Double bond geometry as shown.

L18 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



RM 442561-52-7 CAPLUS  
 CW 1H-Pyrrole-3-carboxamide, 5-[(E)-[4-(3-chlorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-N-(2-(1,1-dioxido-4-thiomorpholinyl)ethyl)-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.



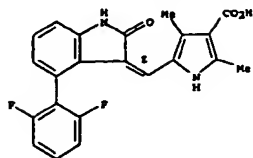
RM 442561-58-2 CAPLUS  
 CW 1H-Pyrrole-3-carboxylic acid, 5-[(E)-[4-(3,6-difluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

Kamal Saeed

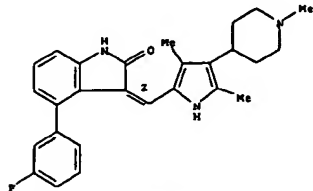
10243942

L18 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



RN 442561-87-7 CAPLUS  
CN 2H-indol-3-one, 3-[[4-(3-fluorophenyl)-1,3-dihydro-2H-indol-3-ylidene]methyl]-4-(3-fluorophenyl)-1,3-dihydro-, [3Z]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.



RN 442561-89-9 CAPLUS  
CN 2H-indol-3-one, 4-(3,5-difluorophenyl)-2-[[4-(3,5-dimethyl-4-(1-methyl-4-piperidinyl)-1H-pyrrol-3-ylidene)methyl]-1,3-dihydro-, [3Z]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L18 ANSWER 5 OF 8 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:31440 CAPLUS

DOCUMENT NUMBER: 136:102386

TITLE: Preparation and use of

4-heteroaryl-3-heteroarylidene-2-indolinones and their use as protein kinase inhibitors

INVENTOR(S): Tang, Peng Cho; Wei, Chung Chen; Huang, Ping; Cui, Jingron

PATENT ASSIGNER(S): Eugen, Inc., USA

SOURCE: PCT Int. Appl., 144 pp.

CODEN: VIXXDJ

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

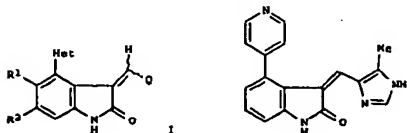
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002002551	A1	20020110	WO 2001-US20768	20010629
W1: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GR, GM, HN, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MY, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AS, BT, BZ, CA, CH, CN, CO, CR, CU, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CP, CO, CI, CM, CA, CN, GW, ML, MR, NG, SN, TD, TG				
US 2002187978	A1	20021212	US 2001-894902	20010629

PRIORITY APPL. INFO.: US 2000-215654P P 20000620

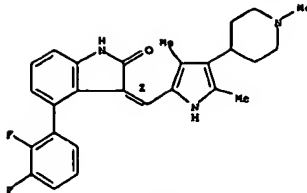
OTHER SOURCE(S): HARPAT 136:102386

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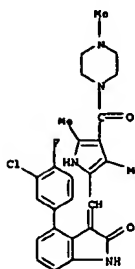


AB Title compds. I (R1-2 = H, alkyl, cycloalkyl, aryl, heteroaryl, heterocyclic, halo, etc.; Het = (un)substituted aro. heterocycle contg. at least one and not more than two N atoms, tetrahydro(thio)pyranyl, (thio)morpholino, piperidinyl, piperazinyl, tetrahydropyridyl, etc.; Q = (un)substituted aro. heterocycle contg. not more than two N atoms, 5-membered ring (un)substituted heterocycle contg. N, O or S, e.g., isoxazolyl, pyrrolyl, indolyl, etc.) with some exceptions, were prepd. Included are 75 synthetic examples and results for several protein tyrosine kinase assays for those compds. For instance, 4-bromoindole was coupled to bis(pinacolato)diborene (DMSO, K2CO3, PdCl3(dppf).bul.OAcCl2, 80.degree.C, 22 h). The resulting dioxaborolane was coupled to 4-bromopyridine.bul.HCl (THF, Pd(PPh3)4, NaOH, 70.degree.C,

L18 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



RN 442562-88-3 CAPLUS  
CN Piperazine, 1-[[5-[[4-(3-chloro-4-(fluorophenyl)-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrol-3-yl]carbonyl]-4-methyl- (9CI) (CA INDEX NAME)



L18 ANSWER 5 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)

6 h) to give the indole which was treated with CSHSH.bul.0r3 (t-BuOH/EtOH/H2O, 1h) followed by aq. (stirred 1 addnl. hour) to give 4-(pyridin-4-yl)-1,1-dihydroindol-2-one as a yellow solid. Condensation of this intermediate with 5-methylimidazole-4-carboxaldehyde (EtOH, piperidine, 2 days) afforded II. II had IC50 = 4.88 nM for FGFR-1 tyrosine kinase and 0.03 nM for cdk2/cyclin A tyrosine kinase. I are useful in treating cancer, immunol. disorders, etc.

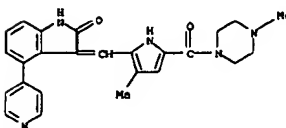
IT 388116-65-2P 388117-30-4P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USBS (Uses)

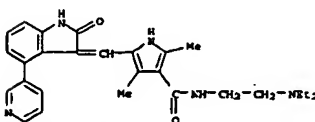
(drug; preps. and use of 4-heteroaryl-3-heteroarylidene-2-indolinones and their use as protein kinase inhibitors)

RN 388116-65-2 CAPLUS

CN Piperazine, 1-[[5-[[1,2-dihydro-2-oxo-4-(4-pyridinyl)-3H-indol-3-ylidene)methyl]-4-methyl-1H-pyrrol-2-yl]carbonyl]-4-methyl- (9CI) (CA INDEX NAME)



RN 388117-30-4 CAPLUS  
CN 1H-Pyrrole-3-carboxamide, N-[2-(diethylanino)ethyl]-5-[[1,2-dihydro-2-oxo-4-(3-pyridinyl)-3H-indol-3-ylidene)methyl]-2,4-dimethyl- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

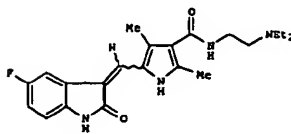
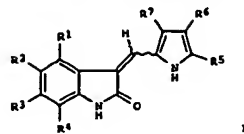
Kamal Saeed

LIS ANSWER 6 OF 8 CAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 3001:617993 CAPLUS  
DOCUMENT NUMBER: 135195497  
TITLE: Preparation of pyrazole substituted 2-indolinone protein kinase inhibitors for treatment of cancer  
(INVENTOR(S)): Tang, Peng Cho; Miller, Todd; Li, Xieoyuan; Suo, Li; Wei, Chung Chou; Shiresian, Shahrsad; Liang, Congxia; Wojkowsky, Tomasz; Wasmatta, Assad S.  
PATENT ASSIGNER(S): Sygen, Inc., USA  
SOURCE: PCT Int. Appl., 225 pp.  
DOCUMNT TYPE: COHER: P1XXD2  
LANGUAGE: Patent  
FAMILY ACC. NUM. COUNT: English 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001060814	A2	20010823	WO 2001-US4813	20010215
WO 2001060814	A3	20020124		
W: AS, AO, AL, AM, AT, AU, AE, BA, BB, BO, BR, BY, BZ, CA, CH, CN, CR, CU, CE, DE, DK, DM, DZ, ES, FI, GB, GD, GE, GH, GM, GW, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LV, MD, MA, MD, MG, MK, MN, MW, MX, ME, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VW, YU, ZA, ZW, AH, AE, BT, BG, BM, BR, BU, BV, CA, CH, CY, CZ, DE, ES, FI, FR, GB, GR, GN, IE, IT, LU, MC, ML, PT, SE, TR, BF, BE, CF, CG, CI, CM, CA, GN, GM, GU, MR, NE, SN, TD, TG				
US 200156292	A1	20010204	US 2001-783264	20010215
EP 1255752	A2	20021113	EP 2001-914376	20010215
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
NO 200203831	A	20021015	NO 2002-3831	20020813
PRIORITY APPL. INFO.:				
			US 2000-183710P	P 200060215
			US 2000-216432P	P 200007005
			US 2000-345322P	P 200010227
			US 2001-US8813	P 20010315

OTHER SOURCE(S): MARPAT 135:195497  
GI

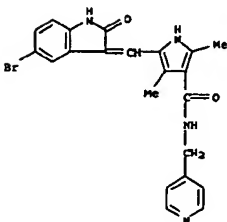
L18 ANSWER 6 OF 8 CAPLIS COPYRIGHT 2003 ACS (Continued)



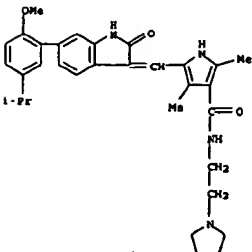
AS The title compo. (I) (wherein R<sup>1</sup> = H, halo, (cyclo)alkyl, (hetero)aryl, (hetero)cyclic, OH, alkoxy, acyl, (un)substituted amino or carbamoyl, etc.; R<sup>2</sup> = H, halo, alkyl, trihalomethyl, OH, alkoxy, CH<sub>2</sub> (hetero)aryl, (un)substituted amino, acyl(amide), or sulfamoyl, etc.; R<sup>3</sup> = H, halo, alkyl, trihalomethyl, OH, alkoxy, (hetero)aryl, (un)substituted acyl, (acyl)amino, sulfamoyl, or alkylsulfonyl, etc.; R<sup>4</sup> = H, halo, alkyl, OH, alkoxy, or (un)substituted amino; R<sup>5</sup> and R<sup>6</sup> = independently M, alkyl, or acyl; R<sup>7</sup> = H, alkyl, (hetero)aryl, or acyl; and their pharmaceutically acceptable salts) were prep'd. as protein kinase modulators for the treatment of cellular disorders such as cancer. For example, 5-fluoro-1,3-dihydroindol-2-one was condensed with 5-formyl-2,4-dimethyl-1H-pyridine-3-carboxylic acid [2-diethylaminoethyl]amide to give II (55%). II was isolated as a crystalline salt. It inhibited protein kinase C- and inhibited PDGF-dependent receptor phosphorylation in cells with an IC<sub>50</sub> value of approx. 0.03 μM. In efficacy expts. against various cancers in mice, II was well tolerated at 80 mg/kg/day, even when dosed continuously for more than 100 days.

IT	342641-67-6P 342641-67-6P 356069-24-4P 356069-42-5P 356069-71-1P
	2L: BAC (Biological) activity or effector, except adverse; BSU
	(Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
	BIOL (Biological study); PREP (Preparation); USES (Uses)
	[prepn. of pyrrole substituted 2-indolinone protein kinase inhibitors
	by condensation of 2-hydroxyindolones with formylpyrroles for treatment
	of cancer and other diseases]
RH	342641-67-6 CAPLUS
	1H-Pyrrole-3-carboxamide, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[(4-pyridinylmethyl)- (SC1) (CA INDEX
NAME	

L10 ANSWER 6 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



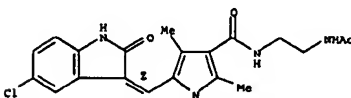
RM 342641-89-8 CAPLUS  
CN 1N-Pyrrole-3-carboxamide, 5-([1,2-dihydro-6-[2-methoxy-5-(1-methylethyl)phenyl]-2-oxo-3H-indol-3-ylidene]methyl)-2,4-dimethyl-N-[2-(1-pyrrolidinyl)ethyl]- (9CI) (CA INDEX NAME)



186069-24-4 CAPLUS  
1H-Pyrrole-3-carboxamide, N-[2-(acetylamino)ethyl]-5-[(2)-(5-chloro-1,3-dihydro-2-oxo-1H-indol-3-ylidene)methyl]-2,4-dimethyl-, (9CI) [CA INDEX NAME]

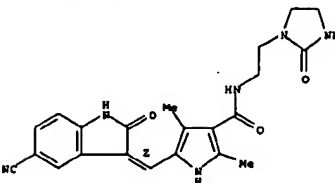
Double bond geometry as shown.

L18 ANSWER 6 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



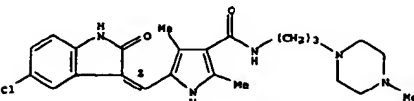
RM 354069-41-5 CAPLAS  
 CN 1H-Pyrrole-3-carboxamide, 5-[[2]-(5-cyano-1,2-dihydro-2-oxo-3H-indol-3-ylidene)ethyl]-2,4-dimethyl-N-(2-(2-oxo-1-imidazolidinyl)ethyl)- (9CI)  
 (CA INDEX NAME)

Double bond geometry as shown.



RN 356069-71-1 CAPLUS  
CN 1H-Pyrrole-3-carboxamide, 5-[(2)-(5-chloro-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[3-(4-methyl-1-piperazinyl)propyl]- (9CI)  
(CA INDEX NAME)

Double bond geometry as shown.



10243942

L18 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:394455 CAPLUS

DOCUMENT NUMBER: 135:19549

TITLE: Preparation of pyrrole substituted 2-indolinones as

antitumor agents

INVENTOR(S): Shenoy, Wamada; Sorauchart, Waramah

PATENT ASSIGNEE(S): Sugen, Inc., USA

SOURCE: PCT Int. Appl., 249 pp.

CODEN: PXXXX2

LANGUAGE: English

DOCUMENT TYPE: Patent

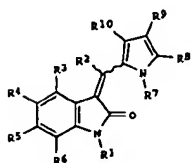
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001037820	A3	20010531	WO 2000-US32277	20001122
WO 2001037820	A3	20011213		
M: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DO, EE, ES, FI, GB, GD, GE, GR, GU, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LE, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TH, TT, TZ, UA, UO, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RN: CH, CM, KE, LE, MW, MZ, SD, SL, SE, TZ, UO, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, DT, BJ, CP, CO, CI, CM, CA, GW, MG, MR, NE, SN, TD, TO				
EP 1331943	A3	20020828	EP 2000-982228	20001122
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, PT, RO, MK, CT, AL, TR				
US 1999-167544P A1 19991124				
WO 2000-US32277 W 20001122				

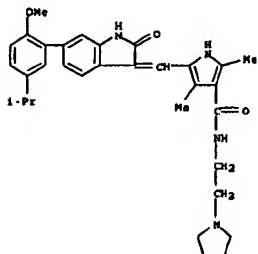
OTHER SOURCE(S): KARPAT 135:19549

Q1



AB The title compds. [I: R1 = H, alkyl, alkenyl, etc.; R2 = H, halo, alkyl, etc.; R3-R6 = H, alkyl, trihaloalkyl, etc.; R7 and R8, R4 and R5, R5 and R6 may combine to form a six membered aryl ring, OCH2O, OCH2CH2O; R7 = H,

L18 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)

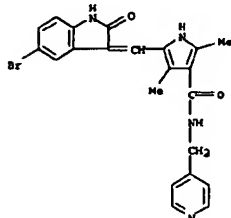


L18 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)  
alkyl, cycloalkyl, etc.; R1-R10 = H, alkyl, trihaloalkyl, etc.) were prep. and formulated. E.g., a multi-step synthesis of I (R1-R7 = H; R8, R10 = Me; R9 = (CH2)2CO2H) which showed 79-86% inhibition of tumor growth of Calu-6 cells in mice at 75 and 100 mg/kg/day, was given. The present invention features formulations of indolinones which compds. are

ionizable as free acids or free bases. The formulation is suitable for parenteral or oral administration, wherein the formulation comprises an ionizable substituted indolinone, and a pharmaceutically acceptable carrier therefor. The term "ionizable substituted indolinone" includes pyrrole substituted 2-indolinones I which, in addn. to being otherwise optionally substituted on both the pyrrole and 2-indolinone portions of the compd., are necessarily substituted on the pyrrole moiety with one or more hydrocarbon chains which themselves are substituted with at least one polar group.

IT 342441-87-EP 342441-89-EP  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prepa. of pyrrole substituted 2-indolinones as antitumor agents)

RN 342441-87-6 CAPLUS  
CH 1H-Pyrrole-3-carboxamide, 5-[(15-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-(4-pyridinylmethyl)- (9CI) (CA INDEX NAME)



RN 342441-89-8 CAPLUS  
CH 1H-Pyrrole-3-carboxamide, 5-[(1,2-dihydro-6-(2-methoxy-5-(1-methylethyl)phenyl)-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-(2-(1-pyrrolidinyl)ethyl)- (9CI) (CA INDEX NAME)

L18 ANSWER 8 OF 8 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2000:660616 CAPLUS

DOCUMENT NUMBER: 134:141324

TITLE: Bioconversion of the anti-angiogenic compound

SUS416

AUTHOR(S): Antonian, Lida; Zhang, Hongbing; Yang, Cheng; Wagner,

Greg; Shawver, Laura K.; Shet, Manjunath; Ogilvie,

Brian; Madan, Ajay; Parkinson, Andrew

CORPORATE SOURCE: SUGEN, Inc., South San Francisco, CA, 94080, USA

SOURCE: Drug Metabolism and Disposition (2000), 28(12),

1505-1512

CODEN: DNGSAL; ISSN: 0090-9556

PUBLISHER: American Society for Pharmacology and Experimental

Therapeutics

DOCUMENT TYPE: Journal

LANGUAGE: English

AB SUS416

[3-(3,5-dimethyl-1H-pyrrol-2-ylmethylene)-1,2-dihydro-indol-2-one],

an inhibitor of VEGF (vascular endothelial growth factor) receptor

tyrosine kinase, Flk-1/KDR (fetal liver kinase 1/kinase insert

domain-contg. receptor), also known as VEGF receptor 2 (V20PR2) is in

advanced clin. trials for treatment of AIDS-related Kaposi's sarcoma and

colorectal and nonsmall cell lung cancers. Since this chem. class has

not

been studied previously with therapeutic intent, the present study was

designed to investigate the in vitro metab. of SUS416 by mouse, rat, dog,

monkey, and human liver microsomes and to identify the major metabolites

of SUS416. An HPLC procedure was developed and validated to resolve and

quantify SUS416 and its metabolites. To evaluate the in vitro metab. of

SUS416, pooled liver microsomes from mice, rats, dogs, monkeys, and

humans

were incubated with SUS416 (25 .mu.M) in the presence of an

NADPH-generating system. In the presence of NADPH, mouse, rat, dog,

monkey, and human liver microsomes converted SUS416 to at least 12, 9, 9,

7, and 6 polar metabolites, resp. Microsomal metab. of SUS416 showed

marked species differences in the levels of different metabolites formed.

The overall rate of SUS416 metab. by liver microsomes from the species

examd. followed the rank order: monkey &gt; mouse .approx. rat &gt; dog

&gt; human. Two major metabolites of SUS416 were identified, a

hydroxymethyl

deriv. of SUS416 (M12) and a carboxylic acid deriv. of SUS416 (M6), by

spectroscopic methods and comparison with authentic compds. Both of

these

oxidative metabolites were further metabolized in vivo through

glucuronidation. The metabolic fate of SUS416 in microsomes from various

species as well as data from in vivo biotransformation in the rat are

discussed.

IT 324047-04-3

RL: BSU (Biological study, unclassified); MPM (Metabolic formation); BIOL

(Biological study); FORM (Formation, nonpreparative)

(biotransformation of anti-angiogenic compd. SUS416)

RN 324047-04-3 CAPLUS

CH .beta.-D-Glucopyranosiduronic acid, [5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-4-methyl-1H-pyrrol-2-yl)methyl (9CI) (CA INDEX NAME)

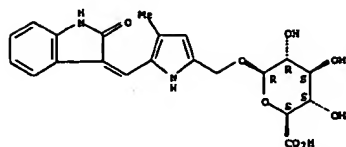
Absolute stereochemistry.

Double bond geometry unknown.

Kamal Saeed

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LIST ANSWER 8 OF 8 CAPLUS COPYRIGHT 2003 ACS (Continued)



REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR  
THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE  
FORMAT

10243942

=> LOGOFF

ALL L# QUERIES AND ANSWER SETS ARE DELETED AT LOGOFF

LOGOFF? (Y)/N/HOLD:Y

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

36.71

784.92

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE

TOTAL

ENTRY

SESSION

CA SUBSCRIBER PRICE

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-5.21

STN INTERNATIONAL LOGOFF AT 11:40:34 ON 12 MAR 2003

Kamal Saeed